

# Implications of modifying the duration requirement of generalized anxiety disorder in developed and developing countries

S. Lee<sup>1\*</sup>, A. Tsang<sup>2</sup>, A. M. Ruscio<sup>3</sup>, J. M. Haro<sup>4</sup>, D. J. Stein<sup>5</sup>, J. Alonso<sup>6</sup>, M. C. Angermeyer<sup>7</sup>,  
E. J. Bromet<sup>8</sup>, K. Demyttenaere<sup>9</sup>, G. de Girolamo<sup>10</sup>, R. de Graaf<sup>11</sup>, O. Gureje<sup>12</sup>, N. Iwata<sup>13</sup>, E. G. Karam<sup>14</sup>,  
J.-P. Lepine<sup>15</sup>, D. Levinson<sup>16</sup>, M. E. Medina-Mora<sup>17</sup>, M. A. Oakley Browne<sup>18</sup>, J. Posada-Villa<sup>19</sup>  
and R. C. Kessler<sup>20</sup>

<sup>1</sup> Department of Psychiatry, The Chinese University of Hong Kong, Hong Kong, PRC; <sup>2</sup> Hong Kong Mood Disorders Center, The Chinese University of Hong Kong, Hong Kong, PRC; <sup>3</sup> Department of Psychology, University of Pennsylvania, Philadelphia, USA; <sup>4</sup> Sant Joan de Deu-SSM, Barcelona, Spain; <sup>5</sup> University of Cape Town, UCT Department of Psychiatry, Cape Town, South Africa; <sup>6</sup> Health Services Research Unit, IMIM-Hospital del mar, CIBER en Epidemiología y Salud Pública (CIBERESP), Barcelona, Spain; <sup>7</sup> Center for Public Mental Health, Gössing a.W, Austria; <sup>8</sup> SUNY Stony Brook, Stony Brook, New York, NY, USA; <sup>9</sup> University Hospital Gasthuisburg, Leuven, Belgium; <sup>10</sup> Regional Health Care Agency, Emilia-Romagna Region, Italy; <sup>11</sup> Netherlands Institute of Mental Health and Addiction, Utrecht, The Netherlands; <sup>12</sup> Department of Psychiatry, University College Hospital, Ibadan, Nigeria; <sup>13</sup> Hiroshima International University, Higashi-Hiroshima, Japan; <sup>14</sup> Institute for Development, Research, Advocacy, and Applied Care (IDRAAC), St. George Hospital University Medical Center, Beirut, Lebanon; <sup>15</sup> Hospital Fernand Widai, Paris, France; <sup>16</sup> Ministry of Health, Mental Health Services, Jerusalem, Israel; <sup>17</sup> National Institute of Psychiatry, Mexico City, Mexico; <sup>18</sup> Department of Rural and Indigenous Health, School of Rural Health, Faculty of Medicine, Nursing and Health Sciences, Monash University, Victoria, Australia; <sup>19</sup> Colegio Mayor de Cundinamarca University, Bogotá, Colombia; <sup>20</sup> Department of Health Care Policy, Harvard Medical School, Boston, USA

**Background.** A number of western studies have suggested that the 6-month duration requirement of generalized anxiety disorder (GAD) does not represent a critical threshold in terms of onset, course, or risk factors of the disorder. No study has examined the consequences of modifying the duration requirement across a wide range of correlates in both developed and developing countries.

**Method.** Population surveys were carried out in seven developing and 10 developed countries using the WHO Composite International Diagnostic Interview (total sample=85 052). Prevalence and correlates of GAD were compared across mutually exclusive GAD subgroups defined by different minimum duration criteria.

**Results.** Lifetime prevalence estimates for GAD lasting 1 month, 3 months, 6 months and 12 months were 7.5%, 5.2%, 4.1% and 3.0% for developed countries and 2.7%, 1.8%, 1.5% and 1.2% for developing countries, respectively. There was little difference between GAD of 6 months' duration and GAD of shorter durations (1–2 months, 3–5 months) in age of onset, symptom severity or persistence, co-morbidity or impairment. GAD lasting ≥12 months was the most severe, persistently symptomatic and impaired subgroup.

**Conclusions.** In both developed and developing countries, the clinical profile of GAD is similar regardless of duration. The DSM-IV 6-month duration criterion excludes a large number of individuals who present with shorter generalized anxiety episodes which may be recurrent, impairing and contributory to treatment-seeking. Future iterations of the DSM and ICD should consider modifying the 6-month duration criterion so as to better capture the diversity of clinically salient anxiety presentations.

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## Introduction

Generalized anxiety disorder (GAD) is increasingly recognized as a prevalent anxiety disorder with characteristic symptoms, significant morbidity and

specific risk factors (WHO, 1993; APA, 2000). Nevertheless, repeated revisions of the GAD diagnostic criteria in recent decades reflect continued uncertainty over the definition and diagnosis of the disorder. Apart from debates over the centrality of pathological worry, the number and type of associated anxiety symptoms, and the level of impairment required for diagnosis (Spitzer & Williams, 1984; Rickels & Rynn, 2001; Slade & Andrews, 2001), a primary focus for

\* Address for correspondence: Professor S. Lee, Director, Hong Kong Mood Disorders Center, 7A, Block E, Staff Quarters, Prince of Wales Hospital, Shatin, NT, Hong Kong.  
(Email: singlee@cuhk.edu.hk)

nosological revision has been the duration criterion of GAD. Although early classifications of pathological anxiety did not specify a minimum duration for diagnosis (Feighner *et al.* 1972; WHO, 1978), efforts to improve diagnostic reliability and differentiation from normal anxiety reactions led to the requirement of a minimum GAD duration of 1 month in the Diagnostic and Statistical Manual of Mental Disorders, 3rd edn (DSM-III, APA, 1980) and an increase to 6 months in the revised DSM-III-R (APA, 1987), the DSM-IV (APA, 2000) and the Diagnostic Criteria for Research of the International Classification of Mental and Behavioral Disorders, 10th edn (ICD-10, WHO, 1993).

Among the anxiety disorders, the requirement of such a long and specific duration is almost unique to GAD, and questions about its necessity remain. This suggests that, despite improved reliability of diagnosis (Brown *et al.* 2001), questions about the validity of GAD are by no means resolved. Following criticisms of the questionable clinical utility of the current duration criterion in defining GAD (Rickels & Rynn, 2001), a number of empirical studies in western countries have recently addressed this question. Taken together, they suggest that GAD lasting 1 month is comparable to GAD lasting  $\geq 6$  months in socio-demographic characteristics, clinical course, pattern of co-morbidity, functional impairment, antecedent childhood adversity, and heritability (Kendler *et al.* 1992, 1994; Bienvenu *et al.* 1998; Maier *et al.* 2000; Carter *et al.* 2001; Hetttema *et al.* 2001; Kessler *et al.* 2005; Angst *et al.* 2006).

Such findings have prompted calls for shortening the duration criterion of GAD in DSM-V (Bienvenu *et al.* 1998; Rickels & Rynn, 2001; Kessler *et al.* 2005; Angst *et al.* 2006; Ruscio *et al.* 2007). Before this change can be considered seriously, however, additional data are needed on at least two fronts. First, the GAD duration criterion originally was increased from 1 to 6 months because of concerns that the shorter duration did not adequately distinguish GAD from normative, transient reactions to stress (Breslau & Davis, 1985). Similar concerns about reduced diagnostic validity and pathologizing of normal stress reactions are likely to be raised for DSM-V. Addressing these concerns will require systematic comparisons of GAD of varying durations on a wide range of relevant validators. Second, although the DSM aspires to be a global diagnostic system, empirical studies of the GAD criteria outside of western countries have been scarce. Recently published data from developing countries such as Nigeria and China have begun to address this gap, but have so far been confined to reporting basic prevalence estimates and sociodemographic correlates of GAD based only on DSM-IV diagnostic criteria (Gureje *et al.* 2006; Shen *et al.* 2006) rather than

evaluating the impact of modifying these criteria. It is therefore desirable to evaluate how varying the duration of GAD may affect its validity in a range of developed and developing countries, ideally using unselected, general population samples to minimize the impact of sampling biases on results.

The present study examined the validity of GAD defined by different durations in a large dataset including representative samples from 10 developed and seven developing countries. After estimating the prevalence of GAD defined by different minimum duration criteria (1 month, 3 months, 6 months, 12 months), we compared the characteristics of four mutually exclusive groups that met the symptom criteria for DSM-IV GAD but differed in the duration of their longest GAD episode.

## Method

### Samples

As part of the World Health Organization's World Mental Health (WMH) Survey Initiative (Demyttenaere *et al.* 2004), 17 countries in the Americas (Colombia, Mexico, USA), Europe (Belgium, France, Germany, Italy, the Netherlands, Spain), Ukraine, the Middle East (Israel, Lebanon), Africa (Nigeria, South Africa) and Asia-Pacific (Japan, New Zealand, People's Republic of China – Beijing, Shanghai) were surveyed. Developing countries were those with a human development index  $< 0.90$ , namely China (Beijing, Shanghai), Colombia, Lebanon, Mexico, Nigeria, South Africa and Ukraine. Developed countries were those with a human development index of  $\geq 0.90$ , namely Belgium, France, Germany, Israel, Italy, Japan, the Netherlands, New Zealand, Spain and the USA (United Nations Development Programme, 2004).

All surveys were based on multi-stage, clustered area probability household samples. Interviews were carried out face to face by trained lay interviewers. The combined total sample size was 85 052 (Table 1). Most of the respondents were aged  $\geq 18$  years, with the exception of respondents from New Zealand (aged  $\geq 16$  years), Japan (aged  $\geq 20$  years) and Israel (aged  $\geq 21$  years). Survey response rates varied, with a weighted average response rate across surveys of 71%. Other than in the Israel survey, where all respondents were administered the full interview, internal subsampling was used to reduce respondent burden by dividing the interview into two parts. Part 1 assessed core mental disorders, including GAD, and was administered to all respondents. Part 2 included additional disorders and correlates relevant to a wide range of survey aims. It was administered to all part 1

respondents who met criteria for any mental disorder as well as a probability sample of other respondents. Part 2 respondents were weighted by the inverse of their probability of selection for part 2 of the interview to adjust for differential sampling. Additional weights were used to adjust for differential probabilities of selection within households and to match the samples to population sociodemographic distributions.

### *Training and field procedures*

The central WMH staff trained bilingual supervisors in each country. Consistent interviewer training documents and standardized translation protocols were used across surveys. The institutional review board of the organization that coordinated the survey in each country approved and monitored compliance with procedures for obtaining informed consent and protecting human subjects.

### *Diagnostic measures*

Mental disorders were assessed using version 3.0 of the World Health Organization Composite International Diagnostic Interview (CIDI) (Kessler & Ustun, 2004), a fully structured lay-administered interview that generates diagnoses according to both ICD-10 (WHO, 1993) and DSM-IV criteria. DSM-IV anxiety, mood and substance-use disorders were included in analyses, as were several disorders that share a common feature of difficulties with impulse control (intermittent explosive disorder, oppositional-deviant disorder, conduct disorder, attention-deficit hyperactivity disorder). Diagnostic hierarchy rules and organic exclusion rules were used in making diagnoses. As detailed elsewhere (Haro *et al.* 2006), blind clinical re-interviews using the Structured Clinical Interview for DSM-IV (SCID, First *et al.* 2002) with a probability subsample of respondents from Spain, Italy, France and the USA found generally good concordance between DSM-IV diagnoses based on the CIDI and the SCID for anxiety (including GAD), mood and substance-use disorders (CIDI diagnoses of impulse-control disorders were not validated).

Respondents were assessed for the symptoms of GAD, and then asked about the number and length of episodes of worry or anxiety experienced in their lifetime and in the last 12 months. 'Episode' was explicitly defined as 'a time lasting 1 month or longer when most days you were (worried or anxious/nervous or anxious/anxious or worried) and also had some of the other problems we just reviewed. The episode ends when you no longer have these feelings for a full month.' Respondents reporting multiple

episodes were grouped for analysis based on their longest lifetime episode.

### *Other measures*

Impairment was assessed by the Sheehan Disability Scale (SDS, Leon *et al.* 1997), which asked respondents to focus on the 1 month in the past year when their GAD was most severe and to rate how much GAD interfered with their home management, work, social life and personal relationships on scales of none (0), mild (1–3), moderate (4–6), severe (7–9) and very severe (10). In addition, role impairment in the past year was assessed by two variables: number of out-of-role days, defined as the number of days out of 365 during which the respondent was 'totally unable' to work or carry out daily activities because of GAD; and role impairment in episode, defined as the percentage of all days in the GAD episode that were spent out of role.

### *Analysis methods*

We estimated the 1-month, 12-month and lifetime prevalence of GAD using the minimum duration requirements of 1, 3, 6 and 12 months. Subsequently, mutually exclusive subgroups with durations of 1–2, 3–5, 6–11 and  $\geq 12$  months were compared on age of onset, clinical course, persistence, role impairment and time to recovery using  $\chi^2$  tests or analysis of variance. Associations with other mental disorders were estimated using a discrete-time survival model with person-year as the unit of analysis, in which variably defined GAD predicted the subsequent first onset of a class of disorders (other anxiety disorders, mood disorders, substance-use disorders, impulse-control disorders). We used the actuarial method (Halli & Vaninadha Rao, 1992) to calculate age-of-onset (AOO) and time to recovery curves for these duration subgroups. Using the Taylor series linearization method (Wolter, 1985) implemented in the SUDAAN software package (Research Triangle Institute, Research Triangle Park, NC, USA), we adjusted for weighting and clustering when calculating standard errors and performing significance tests. Statistical significance was evaluated at the 0.05 level.

## **Results**

### *Prevalence*

The estimated prevalence of GAD increased as the duration criterion was shortened. (Table 2) For developed countries, lifetime prevalence estimates ranged from a low of 3.0% when the minimum duration was 12 months to a high of 7.5% when the minimum

**Table 1.** WMH sample characteristics

Country	Survey	Sample characteristics <sup>a</sup>	Field dates	Age range, years	Sample size			Response rate <sup>c</sup>
					Part 1	Part 2	Part 2 and age ≤ 44 years <sup>b</sup>	
Developed								
Belgium	ESEMeD	Stratified multi-stage clustered probability sample of individuals residing in households from the national register of Belgium residents. NR	2001–2	≥ 18	2419	1043	486	50.6
France	ESEMeD	Stratified multi-stage clustered sample of working telephone numbers merged with a reverse directory (for listed numbers). Initial recruitment was by telephone, with supplemental in-person recruitment in households with listed numbers. NR	2001–2	≥ 18	2894	1436	727	45.9
Germany	ESEMeD	Stratified multi-stage clustered probability sample of individuals from community resident registries. NR	2002–3	≥ 18	3555	1323	621	57.8
Israel	NHS	Stratified multi-stage clustered area probability sample of individuals from a national resident register. NR	2002–4	≥ 21	4859	–	–	72.6
Italy	ESEMeD	Stratified multi-stage clustered probability sample of individuals from municipality resident registries. NR	2001–2	≥ 18	4712	1779	853	71.3
Japan	WMHJ2002–2006	Unclustered two-stage probability sample of individuals residing in households in nine metropolitan areas (Fukiage, Higashi-ichiki, Ichiki, Kushikino, Nagasaki, Okayama, Sano, Tamano, Tendo and Tochigi)	2002–6	≥ 20	3417	1305	425	59.2
Netherlands	ESEMeD	Stratified multi-stage clustered probability sample of individuals residing in households that are listed in municipal postal registries. NR	2002–3	≥ 18	2372	1094	516	56.4
New Zealand	NZMHS	Stratified multi-stage clustered area probability sample of household residents. NR	2004–5	≥ 16	12992	7435	4242	73.3
Spain	ESEMeD	Stratified multi-stage clustered area probability sample of household residents. NR	2001–2	≥ 18	5473	2121	960	78.6
USA	NCS-R	Stratified multi-stage clustered area probability sample of household residents. NR	2002–3	≥ 18	9282	5692	3197	70.9
Developing								
Colombia	NSMH	Stratified multi-stage clustered area probability sample of household residents in all urban areas of the country (approximately 73% of the total national population)	2003	18–65	4426	2381	1731	87.7

Lebanon	LEBANON	Stratified multi-stage clustered area probability sample of household residents. NR	2002–3	≥18	2857	1031	595	70.0
Mexico	M-NCS	Stratified multi-stage clustered area probability sample of household residents in all urban areas of the country (approximately 75% of the total national population)	2001–2	18–65	5782	2362	1736	76.6
Nigeria	NSMHW	Stratified multi-stage clustered area probability sample of households in 21 of the 36 states in the country, representing 57% of the national population. The surveys were conducted in Yoruba, Igbo, Hausa and Efik languages	2002–3	≥18	6752	2143	1203	79.3
People's Republic of China	B-WMH S-WMH	Stratified multi-stage clustered area probability sample of household residents in the Beijing and Shanghai metropolitan areas	2002–3	≥18	5201	1628	570	74.7
South Africa	SASH	Stratified multi-stage clustered area probability sample of household residents. NR	2003–4	≥18	4351	–	–	87.1
Ukraine	CMDPSD	Stratified multi-stage clustered area probability sample of household residents. NR	2002	≥18	4725	1720	541	78.3

WMH, World Mental Health; ESEMeD, The European Study of the Epidemiology of Mental Disorders; NR, nationally representative; NHS, Israel National Health Survey; WMHJ2002–2006, World Mental Health Japan Survey; NZMHS, New Zealand Mental Health Survey; NCS-R, The US National Comorbidity Survey Replication; NSMH, The Colombian National Study of Mental Health; LEBANON, Lebanese Evaluation of the Burden of Ailments and Needs of the Nation; M-NCS, The Mexico National Comorbidity Survey; NSMHW, The Nigerian Survey of Mental Health and Wellbeing; B-WMH, The Beijing World Mental Health Survey; S-WMH, The Shanghai World Mental Health Survey; SASH, South Africa Stress and Health Study; CMDPSD, Comorbid Mental Disorders during Periods of Social Disruption.

<sup>a</sup> Most WMH surveys are based on stratified multi-stage clustered area probability household samples in which samples of areas equivalent to counties or municipalities in the USA were selected in the first stage followed by one or more subsequent stages of geographic sampling (e.g. towns within counties, blocks within towns, households within blocks) to arrive at a sample of households, in each of which a listing of household members was created and one or two people were selected from this listing to be interviewed. No substitution was allowed when the originally sampled household resident could not be interviewed. These household samples were selected from census area data in all countries other than France (where telephone directories were used to select households) and the Netherlands (where postal registries were used to select households). Several WMH surveys (Belgium, Germany and Italy) used municipal resident registries to select respondents without listing households. The Japanese sample is the only totally unclustered sample, with households randomly selected in each of the four sample areas and one random respondent selected in each sample household. Eleven of the seventeen surveys are based on nationally representative household samples, while two others are based on nationally representative household samples in urbanized areas (Colombia, Mexico).

<sup>b</sup> Israel and South Africa did not have an age-restricted part 2 sample. All other countries, with the exception of Nigeria, People's Republic of China and Ukraine (which were age restricted to ≤39 years) were age restricted to ≤44 years.

<sup>c</sup> The response rate is calculated as the ratio of the number of households in which an interview was completed to the number of households originally sampled, excluding from the denominator households known not to be eligible either because of being vacant at the time of initial contact or because the residents were unable to speak the designated languages of the survey.

**Table 2.** Lifetime, 1-year and 1-month prevalence estimates of DSM-IV generalized anxiety disorder when the duration threshold was set at the minimal requirement of 1, 3, 6 and 12 months, in developing and developed countries

	Prevalence of GAD when minimal duration threshold is set at ...			
	1 month	3 months	6 months (DSM-IV criterion)	12 months
<b>Developed countries</b>				
1 month	1.1 (0.1)	0.8 (0.0)	0.7 (0.0)	0.6 (0.0)
12 months	3.2 (0.1)	2.2 (0.1)	1.7 (0.1)	1.3 (0.1)
Lifetime	7.5 (0.2)	5.2 (0.2)	4.1 (0.1)	3.0 (0.1)
<b>Developing countries</b>				
1 month	0.6 (0.1)	0.4 (0.1)	0.4 (0.1)	0.3 (0.0)
12 months	1.4 (0.1)	1.0 (0.1)	0.8 (0.1)	0.7 (0.1)
Lifetime	2.7 (0.1)	1.8 (0.1)	1.5 (0.1)	1.2 (0.1)

DSM-IV, *Diagnostic and Statistical Manual of Mental Disorders*, 4th edn; GAD, generalized anxiety disorder.

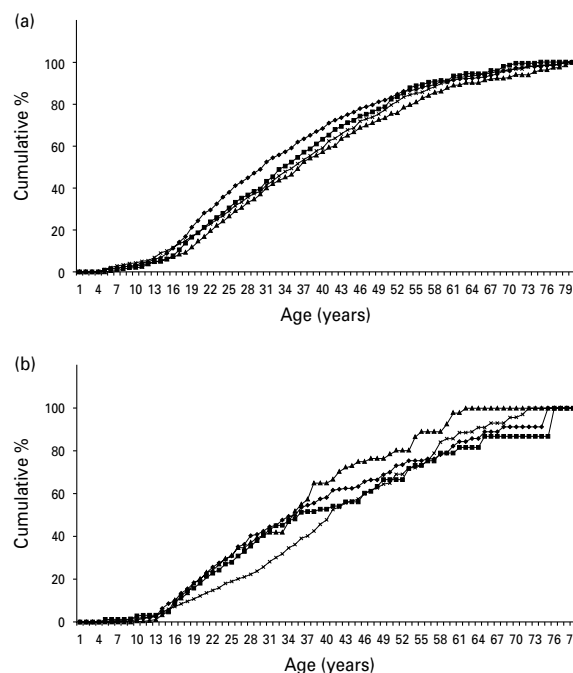
Values are given as percentage (standard error).

duration was 1 month. The corresponding estimates for developing countries were lower, but in the same direction, ranging from 1.2% for 12-month to 2.7% for 1-month minimum duration. The same pattern was evident for 12-month prevalence estimates for developed (1.3–3.2%) and developing (0.7–1.4%) countries as well as for 1-month estimates in both country groups (developed 0.6–1.1%, developing 0.3–0.6%).

### Onset and course

Cumulative AOO distributions were similar in shape for the four mutually exclusive duration subgroups, although the distributions differed significantly [developed  $\chi^2(3)=26.0$ , developing  $\chi^2(3)=18.0$ ,  $p<0.01$ ] (Fig. 1). In both groups of countries, all four subgroups had median AOO in the thirties and rarely had onsets after age 60 years. Mean AOO was also quite similar among the subgroups, falling in the age range of 25–30 years (Table 3). In developed countries, the 1–2 months subgroup had a somewhat later onset than the other three subgroups. In developing countries, there was no significant difference in mean AOO by subgroup.

GAD lasting  $\geq 12$  months exhibited greater persistence than the other subgroups. In both developed and developing countries, this subgroup reported more years with GAD (8.7, 6.5) than the other subgroups, which were more similar to one another (5.3–5.7, 3.5–4.4). Increasing duration of GAD was associated not only with an increase in the longest lifetime GAD



**Fig. 1.** Age of onset of generalized anxiety disorder in (a) developed countries ( $\chi^2=26.0$ ,  $p<0.01$ ) and (b) developing countries ( $\chi^2=18.0$ ,  $p<0.01$ ). —◆—, 1–2 months; —■—, 3–5 months; —▲—, 6–11 months; —×—, 12 months.

episode, but also with more months in episode during the past year. By contrast, the four subgroups did not differ in annual persistence of GAD (the proportion of lifetime cases that had GAD in the past 12 months). The subgroups also did not differ in lifetime persistence of GAD, as they all experienced generalized

**Table 3.** Course of lifetime GAD by specific episode duration (1–2 months, 2–5 months, 6–11 months,  $\geq 12$  months), in developed and developing countries

	Duration of generalized anxiety				<i>F</i> test	
	1–2 months	3–5 months	6–11 months	≥12 months	(df=3)	<i>p</i>
Developed countries						
<i>n</i>	1128	557	604	1603		
Mean age of onset, years (S.E.)	26.7 (0.5)	28.9 (0.7)	30.1 (0.7)	28.9 (0.4)	25.8*	<0.01
Mean number of years with GAD (S.E.)	5.7 (0.3)	5.6 (0.5)	5.3 (0.4)	8.7 (0.3)	55.7*	<0.01
Proportion of years with GAD since onset <sup>a</sup> (S.E.)	0.7 (0.1)	0.7 (0.1)	0.6 (0.1)	0.7 (0)	2.5	0.48
Mean longest duration of GAD, months (S.E.)	1.4 (0)	3.4 (0)	6.8 (0.1)	80.6 (3.9)	457.1*	<0.01
Annual persistence of GAD <sup>b</sup>	41.8 (1.9)	44.2 (2.4)	41.4 (2.3)	42.7 (1.5)	5.5	0.14
Mean number of months in previous year (S.E.) with anxiety among those with GAD in previous year	3.0 (0.2)	3.8 (0.2)	5.2 (0.3)	7.2 (0.2)	272.1*	<0.01
<i>n<sub>p</sub></i>	474	248	251	363		
Developing countries						
<i>n</i>	301	109	108	432		
Mean age of onset, years (S.E.)	25.2 (0.8)	25.8 (1.3)	26.6 (0.7)	29.8 (0.8)	0.5	0.92
Mean number of years with GAD (S.E.)	3.7 (0.3)	3.5 (0.5)	4.4 (0.6)	6.5 (0.5)	14.9*	<0.01
Proportion of years with GAD since onset <sup>a</sup> (S.E.)	0.6 (0.1)	0.6 (0.1)	0.5 (0.1)	0.5 (0)	1.2	0.74
Mean longest duration of GAD, months (S.E.)	1.3 (0)	3.4 (0.1)	6.8 (0.1)	92 (7.5)	148.9*	<0.01
Annual persistence of GAD <sup>b</sup>	53.1 (3.4)	42.6 (5.5)	54.8 (5.4)	56.4 (2.8)	5.4	0.14
Mean number of months in previous year (S.E.) with anxiety among those with GAD in previous year	3.2 (0.3)	3.5 (0.4)	5 (0.5)	6.4 (0.3)	37.1*	<0.01
<i>n<sub>p</sub></i>	157	52	54	251		

GAD, Generalized anxiety disorder; df, degrees of freedom; S.E., standard error; *n<sub>p</sub>*, number of respondents with GAD in previous 12 months.

<sup>a</sup> Number of years in episode divided by the total years between onset age of generalized anxiety and current age.

<sup>b</sup> Percentage of lifetime cases that had generalized anxiety in the past 12 months (ratio of 12-month to lifetime prevalence  $\times 100\%$ ).

\* Significant at  $p < 0.05$ .

anxiety in roughly half of the years since the onset of their respectively defined GAD.

### Severity and impairment

Lifetime GAD severity (mild, moderate, severe) was calculated by submitting 11 nested dichotomous variables representing uncontrollability, distress, and impairment associated with worry and generalized anxiety into an item response theory analysis and trichotomizing the resulting dimension. There was a significant trend in developed [ $\chi^2(6) = 110.5$ ,  $p < 0.01$ ] as well as developing [ $\chi^2(6) = 20.7$ ,  $p < 0.01$ ] countries of fewer mild cases and more severe cases with increasing GAD duration (results not shown, but available on request). The highest proportion of severe cases was in the  $\geq 12$  months subgroup in both developed (41.7%) and developing (36.7%) countries. Nevertheless, severity was substantial even in the 1–2 months subgroup, where the majority of cases (55.9%

in developed and 60.6% in developing countries) were classified as having moderate or severe GAD.

Impairment was also related to duration of GAD (Table 4). In developed countries, the  $\geq 12$  months subgroup was more severely impaired on all SDS domains (4.1–4.7) than the 1–2 months subgroup (3.4–3.8), with the 3–5 and 6–11 months subgroups being intermediate in impairment. Although they differed significantly, the subgroup means for each domain and for the highest-rated domain were within one point of each other on the 0–10 response scale. There was a larger difference between the  $\geq 12$  months subgroup (48.2) and the other subgroups (15.8–24.7) on number of out-of-role days due to GAD. In developing countries, the overall pattern was also that impairment increased with increasing duration, although the increase was less monotonic than the pattern shown in developed countries. This increasing pattern was statistically significant only for social impairment and for days out of role, where the largest difference was

**Table 4.** Past-year impairment in Sheehan Disability Scale domains and days out of role among respondents with lifetime GAD defined by specific episode duration (1–2 months, 2–5 months, 6–11 months,  $\geq 12$  months), in developed and developing countries<sup>a</sup>

	Duration of generalized anxiety				<i>F</i> test, df = 3	<i>p</i>
	1–2 months	3–5 months	6–11 months	≥12 months		
Developed countries						
<i>n<sub>p</sub></i>	474	248	251	363		
Home <sup>b</sup>	3.4 (0.2)	3.6 (0.2)	3.7 (0.2)	4.1 (0.2)	12.0*	0.01
Work <sup>b</sup>	3.7 (0.2)	4.0 (0.3)	3.7 (0.2)	4.3 (0.2)	10.7*	0.01
Relationships <sup>b</sup>	3.8 (0.2)	4.3 (0.3)	4.6 (0.2)	4.7 (0.2)	18.8*	<0.01
Social life <sup>b</sup>	3.7 (0.2)	4.2 (0.3)	4.3 (0.2)	4.4 (0.2)	19.3*	<0.01
Highest-rated domain <sup>b</sup>	5.2 (0.2)	6.1 (0.2)	5.8 (0.2)	6.2 (0.2)	24.5*	<0.01
Days out of role <sup>c</sup>	15.8 (2.7)	24.7 (4.7)	21.3 (4)	48.2 (4.6)	39.9*	<0.01
Role impairment in episode <sup>d</sup> (s.e.)	0.1 (0.0)	0.2 (0.0)	0.1 (0.0)	0.2 (0.0)	13.0*	0.01
Developing countries						
<i>n<sub>p</sub></i>	157	52	54	251		
Home <sup>b</sup>	2.5 (0.3)	3.4 (0.5)	3.4 (0.5)	3.9 (0.3)	7.7	0.05
Work <sup>b</sup>	2.7 (0.3)	3.1 (0.6)	4.0 (0.6)	3.1 (0.3)	5.1	0.16
Relationships <sup>b</sup>	3.0 (0.4)	3.4 (0.5)	3.8 (0.7)	3.0 (0.3)	6.4	0.09
Social life <sup>b</sup>	3.2 (0.4)	3.4 (0.6)	4.6 (0.6)	3.4 (0.3)	8.8*	0.03
Highest-rated domain <sup>b</sup>	4.3 (0.4)	4.9 (0.6)	5.5 (0.5)	4.9 (0.3)	9.0*	0.03
Days out of role <sup>c</sup>	10.8 (2.8)	24.4 (11)	30.8 (10)	29.3 (5.2)	10.3*	0.02
Role impairment in episode <sup>d</sup> (s.e.)	0.1 (0.0)	0.2 (0.1)	0.2 (0.1)	0.1 (0.0)	3.0	0.39

GAD, Generalized anxiety disorder; df, degrees of freedom; *n<sub>p</sub>*, number of respondents with GAD in previous 12 months; S.E., standard error.

<sup>a</sup> Estimated among 12-month cases (*n<sub>p</sub>*).

<sup>b</sup> Mean (S.E.) on the Sheehan Disability Scale, which was rated on a scale of 0 (no impairment), 1–3 (mild), 4–6 (moderate), 7–9 (severe), 10 (very severe impairment).

<sup>c</sup> Mean (S.E.) number of days in the past 12 months during which the respondent was 'totally unable' to work or carry out daily activities because of GAD.

<sup>d</sup> In the past 12 months, proportion of days with GAD that respondents were out of role for the whole day.

\* Significant at  $p < 0.05$ .

between the 1–2 months subgroup (10.8) and the other subgroups (24.4–30.8).

### Co-morbidity and suicidality

All four GAD subgroups were significantly associated with the subsequent first onset of other mental disorders (Table 5). Where significant differences existed, the highest odds of subsequent disorders were found for the  $\geq 12$  month subgroup. Differences between the other subgroups were generally non-monotonic across disorder classes and inconsistent across developed and developing countries. In predicting the onset of any co-morbid mental disorder, only the  $\geq 12$  months subgroup differed significantly from the 1–2 months subgroup in developed countries by exhibiting higher associations with other anxiety disorders and with any mood disorders. None of the subgroups differed significantly in developing countries. In follow-up analyses examining past-year co-morbidity of GAD with individual mental disorders, the four subgroups

had elevated odds ratios (ORs) with every anxiety and mood disorder assessed in the surveys, as well as with intermittent explosive disorder and alcohol abuse (results available on request).

In contrast to GAD of shorter durations, the  $\geq 12$  months subgroup had a significantly elevated risk of suicidality (results not shown, but available on request). The ORs in developed and developing countries were significant for subsequent suicidal ideation (2.0, 2.0), plan (1.7, 2.0) and attempt (1.6, 2.1). The ORs for the other GAD subgroups were non-significant and non-monotonic.

### Treatment and recovery

Duration of GAD in the prior 12-month period was unrelated to 12-month treatment for the disorder (results not shown, but available on request). In developed countries, the lifetime treatment rate for GAD was significantly lower for the 1–2 months subgroup (46.3%) than for the other subgroups (50.7–53.9%),



**Table 5.** Association between generalized anxiety disorder defined with specific duration and risk of subsequent first onset of other disorders, in developed and developing countries<sup>a</sup>

	Duration of generalized anxiety				Wald $\chi^2$ , df = 3	p
	1–2 months	3–5 months	6–11 months	≥ 12 months		
Developed countries						
Any other anxiety disorders <sup>b</sup>	2.4 (1.9–3.0)	2.5 (1.9–3.4)	2.8 (2.0–4.0)	3.7 (3.1–3.4)	13.4*	<0.01
Any mood disorders <sup>c</sup>	2.1 (1.7–2.7)	2.8 (2.0–3.8)	2.6 (1.9–3.6)	3.3 (2.8–4.0)	10.3*	0.02
Any substance-use disorders <sup>b</sup>	2.3 (1.7–3.2)	1.7 (1.0–2.7)	1.5 (0.9–2.6)	2.7 (2.0–3.5)	5.6	0.13
Any impulse-control disorders <sup>d</sup>	4.8 (2.5–9.1)	3.7 (0.9–15.7)	1.4 (0.3–7.4)	2.1 (0.9–5.3)	3.7	0.29
Any disorders <sup>b</sup>	2.1 (1.5–2.9)	1.8 (1.0–3.2)	1.6 (0.9–2.8)	3.4 (2.6–4.5)	10.1*	0.02
Developing countries						
Any other anxiety disorders <sup>b</sup>	3.6 (2.0–6.5)	4.5 (2.1–9.8)	2.9 (1.5–5.4)	6.4 (4.5–9.2)	6.6	0.09
Any mood disorders <sup>c</sup>	3.7 (2.4–5.6)	1.3 (0.6–2.8)	2.5 (1.2–5.3)	4.2 (3.0–6.0)	9.1*	0.02
Any substance-use disorders <sup>b</sup>	2.0 (1.0–4.0)	2.2 (0.7–6.4)	1.9 (0.8–4.4)	3.4 (1.9–6.0)	2.2	0.54
Any impulse-control disorders <sup>d</sup>	1.1 (0.2–7.3)	2.0 (0.5–9.0)	0.8 (0.1–6.3)	11.3 (4.0–31.9)	8.6*	0.04
Any disorders <sup>b</sup>	2.0 (1.0–3.7)	3.9 (1.0–15.5)	4.1 (2.2–7.9)	4.7 (3.0–7.3)	4.9	0.18

df, Degrees of freedom.

Values are given as odds ratio (95% confidence interval).

<sup>a</sup> Based on bivariate analysis using a discrete-time survival model with person-year as the unit of analysis, controlling for person-year, person-year squared, age at interview, sex. No distinctions are made between respondents whose target disorder was active *versus* remitted at the time that the secondary disorder began. Disorders are diagnosed without diagnostic hierarchies.

<sup>b</sup> Weighted on part 2 sample (developed  $n=27\,545$ , developing  $n=15\,579$ ).

<sup>c</sup> Weighted on part 1 sample (developed  $n=50\,791$ , developing  $n=34\,057$ ).

<sup>d</sup> Weighted on part 2 sample in age 44 years or younger (developed  $n=14\,262$ , developing  $n=10\,121$ ).

\* Significant at  $p<0.05$ .

although the proportions were not very different in substantive terms. Duration was not associated with lifetime treatment in developing countries [Wald  $\chi^2(3)=2.7$ ,  $p=0.4$ ], probably because of the low prevalence of GAD and rates of treatment there (18.5–31.4%).

In terms of recovery from GAD, defined as two or more continuous years without symptoms among lifetime cases, the duration subgroups differed from one another significantly for both country groups [developed  $\chi^2(3)=26.0$ , developing  $\chi^2(3)=15.0$ ,  $p<0.01$ ]. A lower proportion of the ≥12 months subgroup recovered than other subgroups in the same period from onset. Regardless of duration, GAD is a chronic disorder, with only half of those affected recovering after about 18–24 years post-onset in developed countries. An even more chronic course was found in developing countries, where the median time to recovery was 24–29 years post-onset. This was most pronounced for the ≥12 months subgroup where the median time to recovery was as much as 50 years after onset. A special finding in developed countries was that the 1–2 and 3–5 months subgroups had a lower proportion of recovery than the 6–11 months subgroup; i.e. the two subgroups took longer to reach the same proportion of

recovery as the 6–11 months subgroup (details of recovery curves not shown, but available on request).

## Discussion

Several limitations of the present study are worthy of note. One is the retrospective assessment of the onset, duration and number of anxiety episodes. Although the probing strategy we used has been shown to improve recall of information such as age of onset (Knauper *et al.* 1999), it was possible that the quality of respondents' recall varied. Recall bias may have been especially likely for respondents with multiple lifetime episodes of varying duration that occurred over a long period of time, or for recollection of more complex diagnostic criteria, such as the number of months when symptoms were present more days than not (Ruscio, 2002). A related limitation is the current lack of consensus among clinicians and researchers about when a GAD episode should be considered to have ended. The operational definition that we used (1 month of full symptom remission) was somewhat arbitrary and may have overestimated duration by requiring a full month without any symptoms rather than a month when symptoms occurred on fewer than

half of the days. Although our use of a fully structured diagnostic instrument and rigorously trained lay interviewers enhanced reliability in the cross-national assessment of mental disorders, the CIDI does not allow symptom clarification and differential diagnosis in the same manner as clinician-administered semi-structured interviews. This might have resulted in inflated associations between GAD and other disorders. While clinical reappraisal studies have suggested reasonably good concordance between CIDI and SCID diagnoses, these studies were performed in only a subset of WMH countries, including China, France, Italy, Spain and the USA (Haro *et al.* 2006; Lee *et al.* 2007b).

With these limitations in mind, the present study showed that the impact of shifting the GAD duration criterion is quite similar across developing and developed countries, though a few differences between the two country groups are worthy of mention. One is the lower prevalence of both lifetime and 12-month GAD in developing countries. This may reflect a generally lower level of psychiatric morbidity (Demyttenaere *et al.* 2004), higher diagnostic thresholds related to respondents having a lower level of mental health literacy, or other methodological factors that lead to a downward bias in the estimation of anxiety disorders in these countries (Shen *et al.* 2006; Chang *et al.* 2008). Levels of GAD-related impairment also differed between developed and developing countries. Respondents with GAD in developing countries reported less impairment on the SDS, despite reporting a similar number of out-of-role days as their counterparts in developed countries. Unlike out-of-role days, which are more objectively defined, there may be cross-cultural differences in appraising and responding to questions about how symptoms have interfered with home management, work, social life and personal relationships. The final difference was the lower treatment rate of GAD in developing countries. This is expected, as high levels of unmet needs for mental health treatment are pervasive in low-income countries (Wang *et al.* 2007).

These differences in prevalence, impairment and treatment notwithstanding, GAD prevalence estimates showed similar increases in both groups of countries as the duration criterion was broadened. Moreover, varying the duration of GAD resulted in very similar changes in onset, course, impairment, comorbidity and recovery rate in developing and developed countries. The findings are consistent with previous western studies (Bienvenu *et al.* 1998; Maier *et al.* 2000; Carter *et al.* 2001; Kessler *et al.* 2005) and extend these studies by providing findings from developing countries that conducted surveys using the same methodology.

Compared with respondents who met the DSM-IV duration requirement of 6 months, respondents with GAD duration of 3–5 months had generally similar age of onset, symptom severity, symptom persistence, role impairment, co-morbidity with other mental disorders, suicidality, treatment pattern and course of recovery. In cases where associations were found between duration and outcomes, differences between the 3–5 and 6–11 months' duration subgroups were either non-monotonic or inconsistent. Moreover, respondents with 3–5 months' GAD recovered more slowly than those with 6–11 months' GAD in developed countries. This is a further indication that GAD of <6 months' duration is not necessarily a milder form of the disorder.

Unlike several western studies that examined durations of 1–6 months and did not find duration to relate to the psychopathological profile of GAD (Bienvenu *et al.* 1998), we found that the  $\geq 12$  months subgroup was more impaired and slower to recover than the other subgroups. In developing countries, where treatment was greatly limited, the median time to recovery was particularly prolonged. Our findings therefore do not completely support the view that duration is of no utility in defining the severity of GAD (Rickels & Rynn, 2001). Because this subgroup represents a chronic form of GAD that is already captured by the DSM-IV 6-month criterion, it is not the focus of the controversy surrounding the duration criterion of GAD. Nevertheless, our findings suggest that episode durations of  $\geq 1$  year may reflect a more severe form of GAD. It is possible that this more chronic form of the disorder may be associated with higher rates of co-morbid axis II pathology or other interpersonal difficulties (Yonkers *et al.* 2000).

Questions related to the duration of GAD are affected by how persistence and termination of episodes are defined. The DSM-IV duration criterion for GAD requires that excessive anxiety and worry occur 'more days than not for at least 6 months' (APA, 2000). If 'more days than not' is interpreted as more than half of the time, the criterion does not specify how the more than 50% of days (i.e. 3 of the 6 months) with anxiety symptoms should be distributed within the 6-month period, nor whether brief periods of complete symptom remission may occur. This is in contrast to the more explicit duration criterion of major depressive episode, in which depressed mood or anhedonia must be present 'most of the day, nearly every day' for 2 weeks (APA, 2000). Studies with clinical samples from specialized treatment centers suggest that most patients with DSM-III-R GAD experience anxiety more days than not (Yonkers *et al.* 1996), but it is unclear whether the same level of persistence would be found in community-based samples. In fact, our data

suggest considerable variability in the course as well as duration of GAD symptoms, with the average respondent in the 1–2 months' duration subgroup reporting a pattern of short episodes of anxiety recurring over many years. However, because subjective perceptions of symptom persistence do not always correspond to the objective number of days with anxiety (Ruscio, 2002), there is a need for prospective research that tracks symptoms on a daily basis for 6 months or more to characterize the course and refine the criteria for GAD.

Our findings have implications for the duration requirement of GAD in DSM-V. If the 6-month duration criterion is maintained, we recommend that DSM-V provide greater description of the longitudinal variability of the course of anxiety. Instead of 'more days than not for at least 6 months', more emphasis can be put on the clinical significance, for example, of individuals who suffer from recurrent short episodes of impairing anxiety.

Shortening the duration criterion in DSM-V – to 1 month, for example – will increase the proportion of people who are diagnosed with GAD in both developed and developing countries and facilitate earlier clinical diagnosis. This modification is supported by our findings that individuals whose symptom duration falls short of the DSM-IV criterion typically suffer moderate to severe symptoms and impairment and have a risk of future mental disorders that is no different than that of respondents who meet the current duration criterion. Although an episode duration of <6 months may be considered subthreshold by DSM-IV standards, affected individuals are not necessarily 'less ill'. Rather, the DSM-IV definition excludes a considerable number of people who suffer from GAD that is <6 months in duration but is nonetheless impairing and recurrent (Kessler *et al.* 2005). This problem is less likely to arise using the Clinical Descriptions and Diagnostic Guidelines of the ICD-10, which specify the duration of GAD as 'variable and usually a few months' (WHO, 1993).

There are, however, potential downsides to reducing the GAD duration requirement. In primary care settings where general practitioners may not have sufficient skills to evaluate pathological worry and quality psychological intervention is barely available, the adoption of a 1-month duration criterion of GAD may contribute to the pathologizing of normal stress responses and indiscriminant pharmacotherapy (Spitzer & Williams, 1984). The inappropriate treatment of normative anxiety responses to stress is especially likely to happen in developing countries, where the quality of mental health services is greatly limited and the chronic use of benzodiazepine tranquillizers is widespread (Lee *et al.* 2007a).

If the duration criterion of GAD is shortened to 1 month in DSM-V, we would recommend a more detailed assessment of pathological anxiety that goes beyond subjective recall of having anxiety 'more days than not' within the 1-month period, so that clinicians can target treatment to those who need it most. Such assessment can be supplemented by the use of dimensional anxiety and impairment scales (Rickels & Rynn, 2001); interviews with collateral informants; and evaluation of concurrent axis II pathology (Yonkers *et al.* 2000), early-onset specific phobias, prior anxiety episodes (especially those lasting  $\geq 12$  months), and other predictors of onset and persistence of psychopathology (Kessler *et al.* 2002). When thus identified, GAD based on a 1-month duration criterion should not be equated with a diagnosis of adjustment disorder, which is often dismissed by clinicians as having no need for treatment. Instead, it should be considered a clinically significant condition which may be especially suitable for early intervention and prevention of secondary morbidity (Ruscio *et al.* 2007). Future trials should examine whether existing therapies would be cost effective in treating 1-month GAD and preventing its recurrence and progression to chronicity (Heuzenroeder *et al.* 2004). Although it is important to recognize the large inter-individual variability in stress responses and to assess the treatment needs of individual patients with care, a shortened GAD duration criterion may not lead to over-treatment. In fact, although major depressive episode has only a 2-week duration criterion, cross-national community epidemiological surveys have shown a large treatment gap for depression across the globe. This gap is especially enormous in developing countries and argues against the inevitability of medicalization in the community (Wang *et al.* 2007).

Finally, it must be emphasized that the present study by no means clarifies all pressing issues related to the diagnostic validity of GAD. Questions about how pathological anxiety should be defined (e.g. its excessiveness, diffuseness, controllability and repetitiveness) and the number and types of associated symptoms that should be required for diagnosis (e.g. somatic symptoms) remain to be examined in future research.

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### Declaration of Interest

Dr Kessler has been a consultant for GlaxoSmithKline Inc., Kaiser Permanente, Pfizer Inc., Sanofi-Aventis, Shire Pharmaceuticals, and Wyeth-Ayerst; has served on advisory boards for Eli Lilly & Company and Wyeth-Ayerst; and has had research support for his epidemiological studies from Bristol-Myers Squibb, Eli Lilly & Company, GlaxoSmithKline, Johnson & Johnson Pharmaceuticals, Ortho-McNeil Pharmaceuticals Inc., Pfizer Inc., and Sanofi-Aventis. Dr Stein has received research grants and/or consultancy honoraria from AstraZeneca, Eli-Lilly, GlaxoSmithKline, Jazz Pharmaceuticals, Johnson & Johnson, Lundbeck, Orion, Pfizer, Pharmacia, Roche, Servier, Solvay, Sumitomo, Tikvah, and Wyeth.

### References

- Angst J, Gamma A, Joseph Bienvenu O, Eaton WW, Ajdacic V, Eich D, Rössler W (2006). Varying temporal criteria for generalized anxiety disorder: prevalence and clinical characteristics in a young age cohort. *Psychological Medicine* **36**, 1283–1292.
- APA (1980). *Diagnostic and Statistical Manual of Mental Disorders*, 3rd edn. American Psychiatric Association: Washington, DC.
- APA (1987). *Diagnostic and Statistical Manual of Mental Disorders*, 3rd edn, revised. American Psychiatric Association: Washington, DC.
- APA (2000). *Diagnostic and Statistical Manual of Mental Disorders*, 4th edn, text revision. American Psychiatric Association: Washington, DC.
- Bienvenu OJ, Nestadt G, Eaton WW (1998). Characterizing generalized anxiety: temporal and symptomatic thresholds. *Journal of Nervous and Mental Disease* **186**, 51–56.

- Breslau N, Davis GC (1985). DSM-III generalized anxiety disorder: an empirical investigation of more stringent criteria. *Psychiatry Research* **15**, 231–238.
- Brown TA, Di Nardo PA, Lehman CL, Campbell LA (2001). Reliability of DSM-IV anxiety and mood disorders: implications for the classification of emotional disorders. *Journal of Abnormal Psychology* **110**, 49–58.
- Carter RM, Wittchen HU, Pfister H, Kessler RC (2001). One-year prevalence of subthreshold and threshold DSM-IV generalized anxiety disorder in a nationally representative sample. *Depression and Anxiety* **13**, 78–88.
- Chang SM, Hahm BJ, Lee JY, Shin MS, Jeon HJ, Hong JP, Lee HB, Lee DW, Cho MJ (2008). Cross-national difference in the prevalence of depression caused by the diagnostic threshold. *Journal of Affective Disorders* **106**, 159–167.
- Demyttenaere K, Bruffaerts R, Posada-Villa J, Gasquet I, Kovess V, Lepine JP, Angermeyer MC, Bernert S, de Girolamo G, Morosini P, Polidori G, Kikkawa T, Kawakami N, Ono Y, Takeshima T, Uda H, Karam EG, Fayyad JA, Karam AN, Mneimneh ZN, Medina-Mora ME, Borges G, Lara C, de Graaf R, Ormel J, Gureje O, Shen Y, Huang Y, Zhang M, Alonso J, Haro JM, Vilagut G, Bromet EJ, Gluzman S, Webb C, Kessler RC, Merikangas KR, Anthony JC, Von Korff MR, Wang PS, Brugha TS, Aguilar-Gaxiola S, Lee S, Heeringa S, Pennell BE, Zaslavsky AM, Ustun TB, Chatterji S, WHO World Mental Health Survey Consortium (2004). Prevalence, severity, and unmet need for treatment of mental disorders in the World Health Organization World Mental Health Surveys. *Journal of the American Medical Association* **291**, 2581–2590.
- Feighner JP, Robins E, Guze SB, Woodruff Jr. RA, Winokur G, Munoz R (1972). Diagnostic criteria for use in psychiatric research. *Archives of General Psychiatry* **26**, 57–63.
- First MB, Spitzer RL, Gibbon M, Williams JBW (2002). *Structured Clinical Interview for DSM-IV Axis I Disorders, Research Version, Non-patient Edition (SCID-I/NP)*. Biometrics Research, New York State Psychiatric Institute: New York.
- Gureje O, Lasebikan VO, Kola L, Makanjuola VA (2006). Lifetime and 12-month prevalence of mental disorders in the Nigerian Survey of Mental Health and Well-Being. *British Journal of Psychiatry* **188**, 465–471.
- Halli SS, Vaninadha Rao K (1992). *Advanced Techniques of Population Analysis*. Plenum Press: New York.
- Haro JM, Arbabzadeh-Bouchez S, Brugha TS, de Girolamo G, Guyer ME, Jin R, Lepine JP, Mazzi F, Reneses B, Vilagut G, Sampson NA, Kessler RC (2006). Concordance of the Composite International Diagnostic Interview version 3.0 (CIDI 3.0) with standardized clinical assessments in the WHO World Mental Health surveys. *International Journal of Methods in Psychiatric Research* **15**, 167–180.
- Hettema JM, Prescott CA, Kendler KS (2001). A population-based twin study of generalized anxiety disorder in men and women. *Journal of Nervous and Mental Disease* **189**, 413–420.
- Heuzenroeder L, Donnelly M, Haby MM, Mihalopoulos C, Rossell R, Carter R, Andrews G, Vos T (2004). Cost-effectiveness of psychological and pharmacological interventions for generalized anxiety disorder and panic disorder. *Australia and New Zealand Journal of Psychiatry* **38**, 602–612.
- Kessler KS, Neale MC, Kessler RC, Heath AC, Eaves LJ (1992). Generalized anxiety disorder in women. A population-based twin study. *Archives of General Psychiatry* **49**, 267–272.
- Kessler KS, Neale MC, Kessler RC, Heath AC, Eaves LJ (1994). Clinical characteristics of familial generalized anxiety disorder. *Anxiety* **1**, 186–191.
- Kessler RC, Andrade LH, Bijl RV, Offord DR, Demler OV, Stein DJ (2002). The effects of comorbidity on the onset and persistence of generalized anxiety disorder in the ICPE surveys. *Psychological Medicine* **32**, 1213–1225.
- Kessler RC, Brandenburg N, Lane M, Roy-Byrne P, Stang PD, Stein DJ, Wittchen HU (2005). Rethinking the duration requirement for generalized anxiety disorder: evidence from the National Comorbidity Survey Replication. *Psychological Medicine* **35**, 1073–1082.
- Kessler RC, Ustun TB (2004). The World Mental Health (WMH) Survey Initiative Version of the World Health Organization (WHO) Composite International Diagnostic Interview (CIDI). *International Journal of Methods in Psychiatric Research* **13**, 93–121.
- Knauper B, Cannell CF, Schwarz N, Bruce ML, Kessler RC (1999). Improving the accuracy of major depression age of onset reports in the US National Comorbidity Survey. *International Journal of Methods in Psychiatric Research* **8**, 39–48.
- Lee S, Tsang A, Chui H, Kwok K, Cheung E (2007a). A community epidemiological survey of generalized anxiety disorder in Hong Kong. *Community Mental Health Journal* **43**, 305–319.
- Lee S, Tsang A, Zhang MY, Huang YQ, He YL, Liu ZR, Shen YC, Kessler RC (2007b). Lifetime prevalence and inter-cohort variation in DSM-IV disorders in metropolitan China. *Psychological Medicine* **37**, 61–71.
- Leon AC, Olsson M, Portera L, Farber L, Sheehan DV (1997). Assessing psychiatric impairment in primary care with the Sheehan Disability Scale. *International Journal of Psychiatry in Medicine* **27**, 93–105.
- Maier W, Gansicke M, Freyberger HJ, Linz M, Heun R, Lecrubier Y (2000). Generalized anxiety disorder (ICD-10) in primary care from a cross-cultural perspective: a valid diagnostic entity? *Acta Psychiatrica Scandinavica* **101**, 29–36.
- Rickels K, Rynn M (2001). Overview and clinical presentation of generalized anxiety disorder. *Psychiatric Clinics of North America* **24**, 1–17.
- Ruscio AM (2002). Delimiting the boundaries of generalized anxiety disorder: differentiating high worriers with and without GAD. *Journal of Anxiety Disorders* **16**, 377–400.
- Ruscio AM, Chiu WT, Roy-Byrne P, Stang PE, Stein DJ, Wittchen HU, Kessler RC (2007). Broadening the definition of generalized anxiety disorder: effects on prevalence and associations with other disorders in the National Comorbidity Survey Replication. *Journal of Anxiety Disorders* **21**, 662–676.

- Shen YC, Zhang MY, Huang YQ, He YL, Liu ZR, Cheng H, Tsang A, Lee S, Kessler RC (2006). Twelve-month prevalence, severity, and unmet need for treatment of mental disorders in metropolitan China. *Psychological Medicine* **36**, 257–267.
- Slade T, Andrews G (2001). DSM-IV and ICD-10 generalized anxiety disorder: discrepant diagnoses and associated disability. *Social Psychiatry and Psychiatric Epidemiology* **36**, 45–51.
- Spitzer RL, Williams JBW (1984). Diagnostic issues in the DSM-III classification of the anxiety disorders. In *Psychiatry Updates: The American Psychiatric Association Annual Review Volume III* (ed. L. Grinspoon), pp. 392–402. American Psychiatric Press: Washington, DC.
- United Nations Development Programme (2004). *Human Development Report 2004: Cultural Liberty in Today's Diverse World*. United Nations Development Programme: New York.
- Wang PS, Angermeyer MC, Borges G, Bruffaerts R, Tat Chiu W, de Girolamo G, Fayyad J, Gureje O, Haro JM, Huang Y, Kessler RC, Kovess V, Levinson D, Nakane Y, Oakley Brown MA, Ormel JH, Posada-Villa J, Aguilar-Gaxiola S, Alonso J, Lee S, Heeringa S, Pennell BE, Chatterji S, Ustun TB (2007). Delay and failure in treatment seeking after first onset of mental disorders in the World Health Organization's World Mental Health Survey Initiative. *World Psychiatry* **6**, 177–185.
- Wolter KM (1985). *Introduction to Variance Estimation*. Springer-Verlag: New York.
- WHO (1978). *Mental Disorders: Glossary and Guide to their Classification in Accordance with the Ninth Revision of the International Classification of Diseases*. World Health Organization: Geneva.
- WHO (1993). *The ICD-10 Classification of Mental and Behavioural Disorders*. World Health Organization: Geneva.
- Yonkers KA, Dyck IR, Warshaw M, Keller MB (2000). Factors predicting the clinical course of generalised anxiety disorder. *British Journal of Psychiatry* **176**, 544–549.
- Yonkers KA, Warshaw MG, Massion AO, Keller MB (1996). Phenomenology and course of generalised anxiety disorder. *British Journal of Psychiatry* **168**, 308–313.